CEP course on Deep Learning for Natural Language Processing

Text Mining in Biomedical and Healthcare Domain

Dr. Shweta Yadav Postdoctoral Research Fellow Wright State University, USA <u>http://shwetanlp.github.io/</u>

Overview



Biomedical Text

INTRODUCTION: Types of Biomedical Texts

Types Of Biomedical Text and its Sources



BIOMEDICAL LITERATURE

MEDLINE

MEDLINE is a bibliographic database of life sciences and biomedical information. It includes bibliographic information for articles from academic journals covering medicine, nursing, pharmacy, dentistry, veterinary medicine, and health care. Wikipedia

History: 1879-present

No. of records: Over 26 million

Record depth: NLM Medical subject headings, abstracts, indexing

Cost: Free

Temporal coverage: 1946-present

Owner: Lister Hill National Center for Biomedical Communications

Image credit: Wikipedia

Producer	U.S. National Library of Medicine (United States)
History	1879-present
Languages	40 languages for current journals, 60 for older journals
	Coverage
Record	NLM Medical subject
depth	headings, abstracts, indexing,
Format	Mostly academic journals; a
coverage	small number of newspapers, magazines, and newsletters; over 40% are for cited articles published in the U.S., about 93% are published in English
Temporal coverage	1946-present
No. of records	Over 26 million
Update	Daily; 2,000-4,000
frequency	references per update
	Links
• Website	

ELECTRONIC MEDICAL RECORDS

John B. Costello, M.D.

2821 North Ballas Rd Ste 165	Phone: 314-995-9988
St Louis, MO 63131	Fax: 314-995-7241

Wednesday, May 05, 2004

Patient

Mr. Steve A Pal 100 Anystreet Saint Louis, MO 63146 44 year old Male DOB: 01/01/60

Visit Date

Thursday, April 22, 2004

Chief Complaint

Sore Throat

History of Present Illness

Injection site identified and marked. Sterile pred with alcohol x3 and ethyl chloride. LOCATION: throat QUALITY: scratchy SEVERITY: moderate DURATION: 2 days

Diagnosis - Major Classic Heartburn 90901

Medications - Current

ACCUPRIL 20 mg 1 1/d Per Oral Accolate 20 mg 1 2/d Per Oral ACTONEL 30 mg 1 1/d Per Oral

Image credit: Wikipedia

SOCIAL MEDIA





I'm loving the new music I've been working on. 6 months off meds I can feel me again. Remember when dark fantasy came out I used to tweet a storm also.



alex joanne @alexvelours - 10h

ignore my greasy eyebrows i just put castor oil on Imao but why is my eyelid like that ong it's swollen and lumpy.. i'm like 60% sure it's an eyelash glue **allergy** but i can't tell HDKDH





lizzie_18070

Hi all,

ive been suffering pretty bad the last few weeks with my <u>anxiety</u> and this week have felt very dizzy ans lightheaded. is this common for anxiety? Or is it possibly something else? does anyone feel this way when their anxiety is up?

0 likes, 9 replies

APPLICATION









Current Medicine

One Treatment Fits All



Future Medicine More Personalized Diagnostics



How to maintain the unstructured biomedical and clinical information ??

DOMAIN KNOWLEDGE BASE



Image credit: [31]

Total # Citations vs. Year of Publication



Year of Publication



1 Billion projected healthrelated apps downloaded a year by 2016

Image credit: [32]

\$500 Billion avoidable annual costs by improving medicine adherence **4X** people over 60 unable to care for themselves by 2050 Exogenous data (Behavior, Socio-economic, Environmental, ...) 60% of determinants of health *Volume, Variety, Velocity, Veracity*

Genomics data 30% of determinants of health Volume

Clinical data 10% of determinants of health Variety Generated per lifetime

6 TB Per lifetime

0.4 TB Per lifetime







Entity Extraction

Patient Data De-Identification (Electronic Medical Records)





Motivation

- Automatically augmenting patient databases
- Unavailability of clinical records for research (even for de-identification) without being de-identified

Challenges

- Inter PHI ambiguity: PHI terms overlap with the non-PHI terms.
 Brown (Doctor name) vs. brown (non-PHI)
- Intra PHI ambiguity: One candidate word seems to belong to two or many different PHI terms.

August (Patient name) vs. August (Date)

- Lexical Variation: For example, variation of the entities such as the '50 yo m', '50 yo M', '55 YO MALE'
- Terminological variation and irregularities: For example '3041023MARY' is the combination of two different PHI categories '3041023' which represents the MEDICALRECORD and 'MARY' which is another PHI category

System	Algorithm	Lexical	Syntactic	Semantics
Guo et al.[7]	SVM	Word , Capitalization, Prefixes/Suffixes, Word Length, Numbers, Regular Expression	POS(Word)	Entity Extract by ANNIE(Doc, Hosp, Loc)
Szarvas et al.[8]	Decision Tree	Word Length, Capitalization, Numbers, Regular Expression, Token Frequency	None	Dictionary Terms (Names , US Loc, Counties, cities, Diseases, Non PHI), Section Headings
Uzuner et al. [9]	SVM	Word , Lexical Bigrams, Capitalization, Punctuation, Numbers, Word Length	POS(Word+2 surrounding)	MeSH ID, dictionary Terms(Names, US and word locations, hospital name)
Wellner et al. [10]	CRF	Word Unigram/Bigram, Surroundings word, Prefixes/Suffixes, capitalization, Numbers, Regular Expression	None	Dictionary Terms (US states, months, General English Terms)
Aramaki et al. [11]	CRF	Word, Surroundings words, capitalization, Word Length, Regular Exp, Sentence Position & Length	POS(Word+2 surroundings word)	Dictionary Terms (Names and Loc)



Feature Engineering

- Bag-of-words
- Part-of-speech (POS) tags
- POS tag of current and surrounding token
- Contextual features
- Sentence information
- Affixes
- Orthographic features
- Word shapes
- Section information
- Task specific features

Dataset (i2b2 2014)

PHI Category	Train	Validation	Test
DOCTOR	2262	183	236
PATIENT	707	28	59
HOSPITAL	1342	141	164
DATE	4154	377	498
LOCATION	93	14	19
PHONE	153	12	13
ID	3200	233	264

PROPOSED APPROACH (Elman RNN)



$$P(y(t) = i | C_m(x_{t-m}^{t+m})) = g(Uh^{(H)}(t) + c)$$
$$g(z_m) = \frac{e^{z_m}}{\sum_{j=1}^{j=k} e^{z_j}}$$
$$h^{(1)}(t) = f(W^{(1)}C_m(x_{t-m}^{t+m}) + V^{(1)}h^{(1)}(t-1) + b)$$
$$h^{(H)}(t) = f(W^{(H)}h^{(H-1)}(t) + V^{(H)}h^{(H)}(t-1) + b)$$

$$C_m(x_{i-m}^{i+m}) = v_{i-m} \oplus \ldots v_i \ldots \oplus v_{i+m}$$

PROPOSED APPROACH (Jordan RNN)



$$h(t) = f(WC_m(x_{t-m}^{t+m}) + VP(y(t-1)) + b)$$

PHI Category	CRF Baseline	Elman RNN	Jordan RNN
PATIENT	58.95	88.89	91.30
DOCTOR	79.08	83.26	85.84
HOSPITAL	60.39	78.03	76.41
LOCATION	55.56	47.83	61.90
PHONE	78.26	88.00	80.00
ID	74.44	90.31	91.68
DATE	94.69	96.74	96.83
Overall	81.39	89.22	90.18



Results with PSO



PHI Category	CRF	CRF+PSO	Elman	Jordan
PATIENT	58.95	59.26	88.89	91.30
DOCTOR	79.08	81.02	83.26	85.84
HOSPITAL	60.39	62.51	78.03	76.41
LOCATION	55.56	55.13	47.83	61.90
PHONE	78.26	78.89	88.00	80.00
ID	74.44	75.41	90.31	91.66
DATE	94.69	95.14	96.74	96.83
OVERALL	81.39	82.58	89.22	90.18



Medical Sentiment Analysis

Social-media Texts (Medical Blogs)

The Digital Patient



Ref. Pew Internet Research. Social Media Usage: 2005-2015. Pew Research Center, October 2015.; http://www.cdwcommunit.com/perspectives/expert-perspectives/todays-digital-patient/

Sample Medical Blog-post



Who is Talking?



Why Not??? Sentiment Analysis



Problem Statement

To prioritize user blog post over two medical sentiment aspects:

- 1. Status of health condition
- 2. Outcome of treatment
Medical Condition



Medication



Proposed Approach (Single Task Learning)



Results

	Task 1: Medical Condition			Task 2: Medication		
Models	Precision	Recall	F-Score	Precision	Recall	F-Score
Baseline 1: SVM	0.42	0.49	0.43	0.74	0.76	0.75
Baseline 2: Random Forest	0.45	0.48	0.46	0.72	0.73	0.73
Baseline 3: MLP	0.41	0.43	0.46	0.74	0.75	0.74
Proposed Approach (CNN)	0.68	0.60	0.63	0.86	0.77	0.82

Method 2: Multi-task Learning

Multi-tasking in NLP



Image Credit:[35]

MTL methods for Deep Learning

Hard parameter sharing

Soft parameter sharing



Image Credit:[36]

Benefits of MTL

- Regularization: it reduces the risk of overfitting as well as the Rademacher complexity of the model
- Representation bias: prefer representations that other tasks also prefer. This will also help the model to generalize to new tasks in the future
- Attention focusing: focus its attention on those features that actually matter as other tasks will provide additional evidence for the relevance or irrelevance of those features.

Feature Space





Shared Private Model

Goal

Adversarial Learning



Adversarial Net Framework



Proposed Approach (Multi Task Learning)



Results

	Task 1: Medical Condition			Task 2: Medication		
Models	Precision	Recall	F-Score	Precision	Recall	F-Score
Baseline 1: MT-LSTM	63.40	61.38	62.37	88.23	77.38	82.45
Baseline 2: ST-LSTM	63.19	62.47	62.83	85.94	77.46	81.48
Proposed Approach	66.82	63.61	65.18	85.83	81.79	83.76

Error Analysis



Approach-3



Result

Models	Techniques Used	Medical Condition			Medications		
		Precision	Recall	F-Score	Precision	Recall	F-Score
Baseline 1	LSTM	65.59	61.23	63.33	85.82	76.19	80.71
Baseline 2	CNN	66.29	61.79	63.96	86.61	76.95	81.49
Baseline 3	MT-LSTM	66.71	64.33	65.5	85.33	81.90	83.58
Proposed Approach	CNN + NLU Features	71.57	67.61	69.53	89.57	86.94	88.23

Disease wise Analysis (Medical Condition)



F1: 'CNN+Emotion (coarse) ', F2: 'CNN+Emotion (fine)', F3: 'CNN+Sentiment word feature', F4: 'CNN+Textual Content Feature', F5: 'Personality', F6: 'Sarcasm'

Disease wise Analysis (Medication)



F1: 'CNN+Emotion (coarse) ', F2: 'CNN+Emotion (fine)', F3: 'CNN+Sentiment word feature', F4: 'CNN+Textual Content Feature', F5: 'Personality', F6: 'Sarcasm'



Pharmacovigilance Mining

Social-media Texts (Medical Blogs)

Introduction

- Medicines: is the applied science or practice of the diagnosis, treatment, and prevention of disease.
- Most medicines have both good and bad effects.
- Bad effects called Adverse Drug Reactions (ADRs), it differs from side effects.
- Side effects whether therapeutic or adverse



ADRs cause over 700,000 emergency department visits each year in the United States

Example of ADRs and side effects

Desired and undesired effects of an aspirin therapy

reduce your headache or fever

reduce the ability of your blood to clot

ASPIRIN® Laters are determined as

× bleeding of intestine

Slide credit: [33]

Pharmacovigilance (PhV)

- Pharmacovigilance (PhV) is the science that concerns with the detection, assessment, understanding and prevention of ADRs
- A Pharmacovigilance (PhV)=drug safety surveillance
- Surveillance for premarketing (i.e. Data from preclinical & clinical trials) and postmarketing (i.e. throughout a drug's market life)

Phv trend to **link** the Preclinical human safety with information from post marketing.



How it Begin??

"Secrets of Seroxat"

BBC Documentary: Panorama broadcasted in 2001

50-minute programme about paroxetine

Crowd Opinion

The programme attracted a record response, including some

65,000 : telephone calls

124,000 : website hits

1,374: emails

FIRST STUDY EXPLORING CROWD INTELLIGENCE

Paroxetine, Panorama and user reporting of ADRs: Consumer intelligence matters in clinical practice and post-marketing drug surveillance

Medawar, C., Herxheimer, A., Bell, A., & Jofre, S. (2002). Paroxetine, Panorama and user reporting of ADRs: Consumer intelligence matters in clinical practice and post-marketing drug surveillance. *International Journal of Risk & Safety in Medicine*, *15*(3, 4), 161-169.

Crowd Intelligence Matters !!

• "Dr Healy confirmed what I already knew. My husband shot himself after 4 days on Seroxat never having been suicidal in his life..."

"I took Seroxat 2 years ago because I have a breathing condition called 'chronic hyperventilation syndrome' which is exacerbated by stress and anxiety. I have never been depressed or had suicidal feelings. However I was prescribed Seroxat to reduce stress & anxiety. A day or two after taking the pills I (went) into a severe state of mental turmoil. I felt really suicidal. It was so severe that all I did was stay in bed for two or three days. Fortunately I recognised Seroxat and stopped taking it immediately."



Text 1: took one pill and 20 minute later had intense pelvic and back pain felt like a miscarriage (i had 3 of them) this intense , horrid pain lasted 1.5 hour then i had spotting and terrible bloating and nausea

Text 2: a 14-year-old girl with newly diagnosed sle developed a pruritic bullous eruption while on prednisone

Text 3: cymbalta , you're driving me insane

Text 4: i have got to stop taking my vyvanse so late !! nosleep add problems

Proposed Approach



Model	Twitter	CADEC	MEDLINE
ST-BLSTM	57.3	51.1	71.91
ST-CNN	67.1	42.0	70.17
CRNN (Huynh et al.,2016)	64.9	48.2	75.5
RCNN (Huynh et al.,2016)	63.6	43.6	74.0
MT-BLSTM (Chowdhury et al.,2016)	63.19	57.62	74.0
MT-BLSTM-Attention (Chowdhury et al.,2016)	65.73	58.27	77.95
Proposed Approach	69.69	65.58	82.18

Conclusion

- Explored the various unstructured form of biomedical text and its application in solving real-world problem.
- Explored deep learning solution based on Elman and Jordan Deep Learning framework for solving patient data de-identification task.
- Exploited the sentiment analysis in medical domain and neural network approach to address the task.
- Explored the unified multi-task learning framework for pharmacovigilance mining that is generic and easily adaptable to extract the pharmacovigilance information from any form of text.

- 1. G. Zhou, J. Zhang, J. Su, D. Shen, and C. Tan, "Recognizing names in biomedical texts: a machine learning approach," Bioinformatics, vol. 20, no. 7, pp. 1178–1190, 2004.
- 2. J.-D. Kim, T. Ohta, Y. Tsuruoka, Y. Tateisi, and N. Collier, "Introduction to the bio-entity recognition task at jnlpba," in Proceedings of the international joint workshop on natural language processing in biomedicine and its applications, pp. 70–75, Association for Computational Linguistics, 2004.
- 3. Park K-M, Kim S-H, Rim H-C, Hwang Y-S (2006) ME-based biomedical named entity recognition using lexical knowledge. ACM Trans Asian Lang Inf Process (TALIP) 5(1):4–21
- 4. B. Settles, "Abner: an open source tool for automatically tagging genes, proteins and other entity names in text," Bioinformatics, vol. 21, no. 14, pp. 3191–3192, 2005.
- Song Y, Kim E, Lee GG, Yi B-k (2004) POSBIOTM-NER in the shared task of BioNLP/NLPBA 2004. In: Proceedings of the international joint workshop on natural language processing in biomedicine and its applications, pp 100–103

- 7. Yikun Guo, Robert Gaizauskas, Ian Roberts, George Demetriou, and Mark Hepple. 2006. Identifying personal health information using support vector machines. In i2b2 workshop on challenges in natural language processing for clinical data, pages 10–11.
- 8. György Szarvas, Richárd Farkas, and András Kocsor. 2006. A multilingual named entity recognition system using boosting and c4. 5 decision tree learning algorithms. In International Conference on Discovery Science, pages 267–278. Springer.
- 9. Özlem Uzuner, Yuan Luo, and Peter Szolovits. 2007. Evaluating the state-of-the-art in automatic de-identification. Journal of the American Medical Informatics Association, 14(5):550–563.
- Ben Wellner, Matt Huyck, Scott Mardis, John Aberdeen, Alex Morgan, Leonid Peshkin, Alex Yeh, Janet Hitzeman, and Lynette Hirschman. 2007. Rapidly retargetable approaches to de-identification in medical records. Journal of the American Medical Informatics Association, 14(5):564–573.
- 11. Eiji Aramaki, Takeshi Imai, Kengo Miyo, and Kazuhiko Ohe. 2006. Automatic deidentification by using sentence features and label consistency. In i2b2 Workshop on Challenges in Natural Language Processing for Clinical Data, pages 10–11.

- M. Krallinger and A. Valencia. Evaluating the detection and ranking of protein interaction relevant articles: the biocreative challenge interaction article sub-task (ias). In Proceedings of the Second Biocreative Challenge Evaluation Workshop, 2007.
- C. Grover, B. Haddow, E. Klein, M. Matthews, L. A. Nielsen, R. Tobin, and X. Wang. Adapting a relation extraction pipeline for the biocreative ii task. In Proceedings of the BioCreAtIvE II Workshop, volume 2, 2007.
- 14. M. Lan, C. L. Tan, and J. Su. Feature generation and representations for protein–protein interaction classification. Journal of biomedical informatics, 42(5):866–872, 2009.
- 15. A. Cohen. Automatically expanded dictionaries with exclusion rules and support vector machine text classifiers: approaches to the biocreative 2 gn and ppi-ias tasks. In Proceedings of the Second BioCreative Challenge Evaluation Workshop, pages 169–174, 2007.
- 16. G. E. Hinton, N. Srivastava, A. Krizhevsky. Improving neural networks by preventing co-adaptation of feature detectors. arXiv preprint arXiv:1207.0580, 2012.
- W. A. Baumgartner Jr, Z. Lu, An integrated approach to concept recognition in biomedical text. In Proceedings of the Second BioCreative Challenge Evaluation Workshop, volume 23, pages 257–71. Centro Nacional de Investigaciones Oncolog-icas (CNIO) Madrid, Spain, 2007

- 18. Hua L., Quan C.**A shortest dependency path based convolutional neural network for protein-protein relation extraction** BioMed. Res. Int., 2016 (2016)
- 19. Choi S.-P.Extraction of protein–protein interactions (ppis) from the literature by deep convolutional neural networks with various feature embeddings J. Inf. Sci. (2016)
- 20. Y. Peng, Z. Lu, Deep learning for extracting protein-protein interactions from biomedical literature, arXiv preprint <u>arXiv:1706.01556</u>.
- 21. Qian L., Zhou G.**Tree kernel-based protein–protein interaction extraction from biomedical** literature J. Biomed. Inform., 45 (3) (2012), pp. 535-543
- 22. Li L., Guo R., Jiang Z., Huang D.**An approach to improve kernel-based protein–protein** interaction extraction by learning from large-scale network data Methods, 83 (2015), pp. 44-50
- 23. Choi S.-P., Myaeng S.-H. **Simplicity is better: revisiting single kernel ppi extraction.** Proceedings of the 23rd International Conference on Computational Linguistics, Association for Computational Linguistics (2010), pp. 206-214

- 24. Nikfarjam, A., & Gonzalez, G. H. (2011). Pattern mining for extraction of mentions of adverse drug reactions from user comments. In *AMIA Annual Symposium Proceedings* (Vol. 2011, p. 1019). American Medical Informatics Association.
- 25. Yates, Andrew, and Nazli Goharian. "ADRTrace: detecting expected and unexpected adverse drug reactions from user reviews on social media sites." In *European Conference on Information Retrieval*, pp. 816-819. Springer, Berlin, Heidelberg, 2013.
- 26. Sarker, A., & Gonzalez, G. (2015). Portable automatic text classification for adverse drug reaction detection via multi-corpus training. *Journal of biomedical informatics*, *53*, 196-207.
- 27. O'Connor, K., Pimpalkhute, P., Nikfarjam, A., Ginn, R., Smith, K. L., & Gonzalez, G. (2014). Pharmacovigilance on twitter? Mining tweets for adverse drug reactions. In *AMIA annual symposium proceedings* (Vol. 2014, p. 924). American Medical Informatics Association.
- 28. Trung Huynh, Yulan He, Alistair Willis, and Stefan Rüger. 2016. Adverse drug reaction classification with deep neural networks. Coling.
- 29. Shaika Chowdhury, Chenwei Zhang, and Philip S. Yu. 2018. Multi-task pharmacovigilance mining from social media posts. In Proceedings of the 2018 World Wide Web Conference, WWW '18, pages 117–126, Republic and Canton of Geneva, Switzerland. International World Wide Web Conferences Steering Committee.

- 30. Simmons, M., Singhal, A., & Lu, Z. (2016). Text mining for precision medicine: bringing structure to EHRs and biomedical literature to understand genes and health. In *Translational Biomedical Informatics* (pp. 139-166). Springer, Singapore.
- 31. Olivier Bodenreider. (2010, October 21). The Unified Medical Language System What is it and how to use it? Retrieved from

https://www.slideshare.net/roger961/the-unified-medical-language-system-what-is-it-and-how-to-use-it.

- 32. Kerrie Holley. (2013). Technological Trend of the Future: Next Era of Computing. Retrieved from https://www.slideshare.net/IBMSMMA/ibm-summit-2013-16-9next-era
- 33. DigiMind. (2016). Pharmacovigilance & Social Media. Retrieved from <u>https://www.slideshare.net/digimind/pharmacovigilance-social-media</u>
- 34. Liu, P., Qiu, X., & Huang, X. (2017). Adversarial multi-task learning for text classification. *arXiv preprint arXiv:1704.05742*.
- 35. Alexandr Honchar. (2018). Multitask learning: teach your AI more to make it better. Retrieved from https://towardsdatascience.com/multitask-learning-teach-your-ai-more-to-make-it-better-dde116c2cd40
- 36. Sebastian Ruder. (2017). An Overview of Multi-Task Learning In Deep Neural Networks. Retrieved from <u>https://ruder.io/multi-task/</u>
- 37. Ian Goodfellow. (2017). Introduction to GANs, NIPS 2016. Retrieved from https://www.youtube.com/watch?v=9JpdAg6uMXs&frags=pl%2Cwn

THANK YOU!